

**PATENT**

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

In re Application of:  
Inderjit Singh

Serial No.: 10/596,198

Filed: June 2, 2006

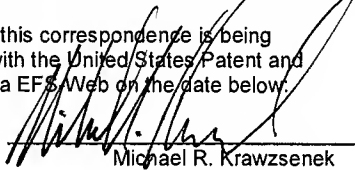
For: METHODS AND COMPOSITIONS FOR  
THE PREVENTION AND TREATMENT  
OF INFLAMMATORY DISEASES OR  
CONDITIONS

Group Art Unit: 2622

Examiner: Unknown

Atty. Dkt. No.: MESC:013US

Confirmation No.: 3351

CERTIFICATE OF ELECTRONIC TRANSMISSION	
I hereby certify that this correspondence is being electronically filed with the United States Patent and Trademark Office via EFS Web on the date below.	
May 10, 2007	
Date	Michael R. Krawzsenek

**INFORMATION DISCLOSURE STATEMENT**

Commissioner for Patents  
P.O. Box 1450  
Alexandria, Virginia 22313-1450

Sir:

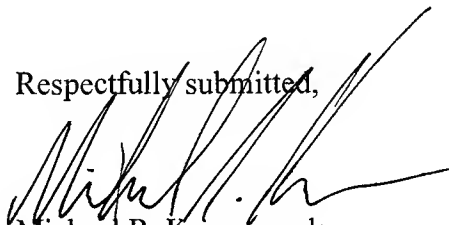
In compliance with the duty of disclosure under 37 C.F.R. § 1.56, it is respectfully requested that this Information Disclosure Statement be entered and the documents listed on attached Form PTO-1449 be considered by the Examiner and made of record. Copies of the listed documents required by 37 C.F.R. § 1.98(a)(2) are enclosed for the convenience of the Examiner.

In accordance with 37 C.F.R. §§ 1.97(g), (h), this Information Disclosure Statement is not to be construed as a representation that a search has been made, and is not to be construed to be an admission that the information cited is, or is considered to be, material to patentability as defined in 37 C.F.R. § 1.56(b).

The present Information Disclosure Statement is being filed prior to the receipt of a first Official Action reflecting an examination on the merits, and hence is believed to be timely filed in accordance with 37 C.F.R. § 1.97(b). No fees are believed to be due in connection with the filing of this Information Disclosure Statement, however, should any fees under 37 C.F.R. §§ 1.16 to 1.21 be deemed necessary for any reason relating to these materials, the Commissioner is authorized to deduct the appropriate fees from Fulbright & Jaworski Deposit Account No.: 50-1212/MESC:013US.

Applicant respectfully requests that the listed documents be made of record in the present case.

Respectfully submitted,



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Date: May 10, 2007

Form PTO-1449 (modified)		Atty. Docket No. <b>MESC:013US</b>	Serial No. <b>10/596,198</b>
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		Filing Date: <b>June 2, 2006</b>	Group: <b>2622</b>
U.S. Patent Documents <i>See Page 1-2</i>	Foreign Patent Documents <i>See Page 2</i>	Other Art <i>See Page 2-29</i>	

### U.S. Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Name	Class	Sub Class	Filing Date of App.
	A1	2001/0041726	11/15/01	Bandarage <i>et al.</i>	514	361	12/22/00
	A2	2003/0104348	06/05/03	Vodovotz <i>et al.</i>	435	1.2	08/28/02
	A3	2007/0027094	02/01/07	Singh	514	43	08/15/05
	A4	5,118,601	06/02/92	Gruber	435	5	08/31/89
	A5	5,130,333	07/14/92	Pan <i>et al.</i>	514	460	10/19/90
	A6	5,286,895	02/15/94	Harris <i>et al.</i>	560	60	12/07/92
	A7	5,498,536	03/12/96	Khandke	435	200	04/22/94
	A8	5,545,625	08/13/96	Gross <i>et al.</i>	514	33	12/12/94
	A9	5,658,889	08/19/97	Gruber <i>et al.</i>	514	43	12/14/94
	A10	5,695,761	12/09/97	Denhardt <i>et al.</i>	424	184.1	12/23/93
	A11	5,912,019	06/15/99	Singh	424	608	10/16/97
	A12	5,965,553	10/12/99	Bell <i>et al.</i>	514	211.09	06/12/97
	A13	5,968,983	10/19/99	Kaesemeyer	514	564	04/10/97
	A14	6,080,778	06/27/00	Yankner <i>et al.</i>	514	451	03/23/98
	A15	6,147,109	11/14/00	Liao <i>et al.</i>	514	460	08/11/98
	A16	6,204,248	03/20/01	Demopoulos <i>et al.</i>	514	21	12/09/99
	A17	6,312,662	11/06/01	Erion <i>et al.</i>	424	9.1	09/08/99
	A18	6,472,390	11/13/01	Stamler <i>et al.</i>	514	236.5	11/13/01
	A19	6,511,800	01/28/03	Singh	435	4	05/25/00
	A20	6,552,075	04/22/03	Gribble <i>et al.</i>	514	522	08/09/01
	A21	6,596,701	07/22/03	Schwartz <i>et al.</i>	514	46	03/16/98
	A22	6,605,729	08/12/03	Byrn <i>et al.</i>	548	537	06/28/02
	A23	6,649,629	11/18/03	Bandarage <i>et al.</i>	514	326	12/22/00

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	A24	6,656,966	12/02/03	Garvey <i>et al.</i>	514	449	06/22/01
	A25	6,673,831	01/06/04	Tolbert	514	252.17	04/27/98
	A26	7,049,058	05/23/06	Singh	435	4	10/18/02

### Foreign Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Country	Language
	B1	WO 93/14749	08/05/93	WIPO	English
	B2	WO 95/06470	03/09/95	WIPO	English
	B3	WO 98/09653	03/12/98	WIPO	French (English Abstract)

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	C1	"Nymox pursuing new approaches to fighting Alzheimer's disease," <i>Business Wire</i> , July 2, 2004.
	C2	"Statins-the next miracle drug? Beyond cholesterol lowering," <i>Clinician Reviews</i> , 14(6):84(3), June 1, 2004.
	C3	Abud-Mendoza <i>et al.</i> , "Therapy with statins in patients with refractory rheumatic diseases: a preliminary study," <i>Lupus</i> , 12(8):607-611, 2003.
	C4	Akiyama <i>et al.</i> , "Cell mediators of inflammation in the Alzheimer disease brainm" <i>Alzheimer Dis. Assoc. Disord.</i> , 1:S47-53, 2000.
	C5	Akiyama <i>et al.</i> , "Inflammation and Alzheimer's disease," <i>Neurobiol. Aging</i> , 21:383-421, 2000.
	C6	Amerongen <i>et al.</i> , "Simvastatin improves disturbed endothelial barrier function," <i>Circulation</i> , 102:2803-2809, 2000.
	C7	Anderson <i>et al.</i> , "Astrocytes and stroke: networking for survival?," <i>Neurochem. Res.</i> , 28:293-305, 2003.

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Exam. Init.	Ref. Des.	Citation
	C8	Andoh <i>et al.</i> , "The role of neuronal nitric oxide synthase (NOS1) and related cyclic GMP-PKG signaling pathway on preconditioning-mediated neuroprotection," <i>Society for Neuroscience Abstracts</i> , 2001.
	C9	Appel <i>et al.</i> , "Lack of interaction between fluvastatin and oral hypoglycemic agents in healthy subjects and in patients with non-insulin-dependent diabetes mellitus," <i>Am. J. Cardiol.</i> , 76:29A-32A, 1995.
	C10	Arai <i>et al.</i> , "Lactosylceramide stimulates human neutrophils to upregulate Mac-1, adhere to endothelium, and generate reactive oxygen metabolites in vitro," <i>Circ. Res.</i> , 82:540-547, 1998.
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	C12	Azuma <i>et al.</i> , "HMG-CoA reductase inhibitor attenuates experimental autoimmune myocarditis through inhibition of T cell activation," <i>Cardiovascular Res.</i> , 64:412-420, 2004.
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	C14	Bal-Price and Brown, "Inflammatory neurodegeneration mediated by nitric oxide from activated glia-inhibiting neuronal respiration, causing glutamate release and excitotoxicity," <i>J. Neurosci.</i> , 21:6480-6491, 2001.
	C15	Barber <i>et al.</i> , "Biochemistry of Ischemic Stroke," <i>Adv. Neuro.</i> , 192:151-164, 2003.
	C16	Barone and Feuerstein, "Inflammatory mediators and stroke: new opportunities for novel therapeutics," <i>J. Cereb. Blood Flow Metab.</i> , 19:819-834, 1999.
	C17	Bauer <i>et al.</i> , "Inflammation in the Nervous System: The Human Perspective," <i>Glia</i> , 36:235-243, 2001.
	C18	Baughman, "Dopamine-transporter density in patients with ADHD," <i>The Lancet</i> , 355:1460-61, 2000.
	C19	Bedoya <i>et al.</i> , "Pyrrolidine dithiocarbamate prevents IL-1-induced nitric oxide synthase mRNA, but not superoxide dismutase mRNA, in insulin producing cells," <i>Biochemical and Biophysical Research Communications</i> , 210:816-822, 1995.
	C20	Bellosta <i>et al.</i> , "Direct vascular effects of HMG-CoA reductase inhibitors," <i>Atheroscl.</i> , 137:S101-109, 1998.
	C21	Bellosta <i>et al.</i> , "Non-lipid related effects of statins," <i>Ann. Med.</i> , 32:164-176, 2000.

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	C22	Benveniste <i>et al.</i> , "Cytokines: influence on glial cell gene expression and function," <i>Chem. Immunol.</i> , 69:31-75, 1997.
	C23	Berisha <i>et al.</i> , "Nitric oxide as a mediator of oxidant lung injury due to paraquat," <i>Proc. Natl. Acad. Sci. USA</i> , 91:7445-7449, 1994.
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	C25	Bhunja <i>et al.</i> , "Lactosylceramide mediates tumor necrosis factor-alpha-induced intercellular adhesion molecule-1 (ICAM-1) expression and the adhesion of neutrophil in human umbilical vein endothelial cells," <i>J. Biol. Chem.</i> , 273:34349-34357, 1998.
	C26	Bhunja <i>et al.</i> , "Redox-regulated signaling by lactosylceramide in the proliferation of human aortic smooth muscle cells," <i>J. Biol. Chem.</i> , 272:15642-15649, 1997.
	C27	Biesenbach and Zazgornik, "Lovastatin in the treatment of hypercholesterolemia in nephrotic syndrome due to diabetic nephropathy state IV-V," <i>Clin. Nephrology</i> , 37:274-279, 1992.
	C28	Blazquez <i>et al.</i> , "The AMP-activated protein kinase prevents ceramide synthesis de novo and apoptosis in astrocytes," <i>FEBS Lett.</i> , 489:149-153, 2001.
	C29	Blight, "Anoxia Physiology of Chronic Spinal Cord Injury in the Cat: Intracellular Recording in Vitro," <i>Neuroscience</i> , 10:1471-1486, 1983.
	C30	Bo <i>et al.</i> , "Induction of nitric oxide synthase in demyelinating regions of multiple sclerosis brains," <i>Ann. Neurol.</i> , 36:778-786, 1994.
	C31	Boese, "Effect of cyclic GMP-dependent vasodilators on the expression of inducible nitric oxide synthase in vascular smooth muscle cells: role of cyclic AMP," <i>Br. J Pharmacology</i> , 119:707-715, 1996.
	C32	Bolanos and Almeida, "Roles of nitric oxide in brain hypoxia-ischemia," <i>Biochem. Biophys. Acta</i> , 1411:415-436, 1999.
	C33	Boughton-Smith <i>et al.</i> , "Nitric oxide synthase activity in ulcerative colitis and Crohn's disease," <i>Lancet</i> , 342:338-340, 1993.
	C34	Bowen and Guyton, "Nonpharmacologic and pharmacologic treatment of patients with low levels of high-density lipoprotein cholesterol," <i>Current Science</i> , 2:58-63, 2000.

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	C35	Bredesen, "Neural apoptosis," <i>Ann. Neurol.</i> , 38:839-851, 1995.
	C36	Brugg <i>et al.</i> , "Ceramide induces apoptosis in cultured mesencephalic neurons," <i>J. Neurochem.</i> , 66:733-739, 1996.
	C37	Brune <i>et al.</i> , "Nitric oxide and its role in apoptosis," <i>Eur. J. Pharmacol.</i> , 351:261-272, 1998.
	C38	Buemi <i>et al.</i> , "Effect of fluvastatin on proteinuria in patients with immunoglobulin A nephropathy," <i>Clin. Phar. &amp; Therapeutics</i> , 67:427-431, 2000.
	C39	Buisson <i>et al.</i> , "The neuroprotective effect of a nitric oxide inhibitor in a rat model of focal cerebral ischaemia," <i>Br. J. Pharmacol.</i> , 106:766-767, 1992.
	C40	Bull <i>et al.</i> , "Nitric Oxide Synthase in the Human Dermal Microvasculature," <i>J. Invest. Derm.</i> , 103:435, 1994 (Abstract No. 231).
	C41	Burney <i>et al.</i> , "The chemistry of DNA damage from nitric oxide and peroxynitrite," <i>Mutat. Res.</i> , 424:37-49, 1999.
	C42	Cartier <i>et al.</i> , "Retroviral-mediated gene transfer corrects very-long-chain fatty acid metabolism in adrenoleukodystrophy fibroblasts," <i>Proc. Natl. Acad. Sci. USA</i> , 92:1674-1678, 1995.
	C43	Casey <i>et al.</i> , "p21ras is modified by a farnesyl isoprenoid," <i>Proc. Natl. Acad. Sci.</i> , 86:8323-27, 1989.
	C44	Catani <i>et al.</i> , "Inhibition of clotting factor XIII activity by nitric oxide," <i>Biochem. Biophys. Res. Commun.</i> , 249:275-278, 1998.
	C45	Cetkovic-Cvrlje <i>et al.</i> , "Nicotinamide and Dexamethasone Inhibit Interleukin-1-Induced Nitric Oxide Production by RINm5F Cells without Decreasing Messenger Ribonucleic Acid Expression for Nitric Oxide Synthase," <i>Endocrinology</i> , 133:1739-1743, 1993.
	C46	Chatterjee, "Sphingolipids in atherosclerosis and vascular biology," <i>Arterioscler. Thromb Vasc. Biol.</i> , 18:1523-1533, 1998.
	C47	Chen and Rosazza, "A bacterial nitric oxide synthase from a Nocardia species," <i>Biochem. Biophys. Res. Commun.</i> , 203:1251-1258, 1994.
	C48	Chieuh <i>et al.</i> , "S-nitrosoglutathione/nitric oxide system: A possible antioxidative cellular defense system in the brain," <i>Jpn. J. Pharmacol.</i> , pp. 8P, 1997 (Abstract No. 22).

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Exam. Init.	Ref. Des.	Citation
	C49	Chiueh and Andoh, "Nitric oxide(cntdotNO) production through transfection and induction of NOS1 increases tolerance to 1-methyl-4-phenylpyridinium (MPP+)-induced neurotoxicity," <i>Society for Neuroscience Abstracts</i> , 2000.
	C50	Chiueh and Rauhala, "The redox pathway of S-nitrosoglutathione, glutathione and nitric oxide in cell to neuron communications," <i>Free Radic. Res.</i> , 31:641-650, 1999.
	C51	Chiueh <i>et al.</i> , "Enhanced Hydroxyl Radical Generation by 2'-Methyl Analog of MPTP: Suppression by Clorgyline and Deprenyl," <i>Synapse</i> , 11:346-348, 1992.
	C52	Chiueh, "Neuroprotective properties of nitric oxide," <i>Anns. NY Acad. Sci.</i> , 890:301-311, 1999 (Abstract Only).
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	C54	Choi and Lipton, "Redox modulation of the NMDA receptor," <i>Cell Mol. Life Sci.</i> , 57:1535-1541, 2000.
	C55	Chung <i>et al.</i> , "Nitric oxide as a bioregulator of apoptosis," <i>Biochem. Biophys. Res. Commun.</i> , 282:1075-1079, 2001.
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	C57	Corbett and McDaniel, "Selective inhibition of inducible nitric oxide synthase by aminoguanidine," <i>Methods Enzymol.</i> , 268:398-408, 1996.
	C58	Corsini <i>et al.</i> , "Non-lipid-related effects of 3-hydroxy-3-methylglutaryl coenzyme A reductase inhibitors," <i>Cardiology</i> , 87:458-468, 1996.
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	C60	Cross <i>et al.</i> , "Aminoguanidine, an inhibitor of inducible nitric oxide synthase, ameliorates experimental autoimmune encephalomyelitis in SJL mice," <i>J. Clin. Invest.</i> , 93:2684-2690, 1994.
	C61	Cui <i>et al.</i> , "Oxidative DNA damage precedes DNA fragmentation after experimental stroke in rat brain," <i>Faseb J.</i> , 14:955-967, 2000.

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	C62	Culmsee <i>et al.</i> , "AMP-activated protein kinase is highly expressed in neurons in the developing rat brain and promotes neuronal survival following glucose deprivation," <i>J. Mol. Neurosci.</i> , 17:45-58, 2001.
	C63	Cuzzocrea <i>et al.</i> , "Beneficial effects of n-acetylcysteine on ischemic brain injury," <i>British Journal of Pharmacology</i> , 130:1219-1226, 2000.
	C64	Davis <i>et al.</i> , "Novel effects of nitric oxide," <i>Annu. Rev. Pharmacol. Toxicol.</i> , 41:203-236, 2001.
	C65	Davoli <i>et al.</i> , "Immunohistochemical and Biochemical Assessment of Caspase-3 Activation and DNA Fragmentation following Transient Focal Ischemia in the Rat," <i>Neuroscience</i> , 115:125-136, 2002.
	C66	Dawson <i>et al.</i> , "Nitric oxide as a mediator of neurotoxicity," <i>NIDA Res. Monogr.</i> , 136:258-271, 1993.
	C67	Devlin <i>et al.</i> , "Nitric oxide generation. A predictive parameter of acute allograft rejection," <i>Transplantation</i> , 58:592-595, 1994.
	C68	Dignam <i>et al.</i> , "Eukaryotic gene transcription with purified components," <i>Methods Enzymol.</i> , 101:582-598, 1983.
	C69	Ding-Zhou <i>et al.</i> , "L-NAME reduces infarction, neurological deficit and blood-brain brain disruption following cerebral ischemia in mice," <i>Eur. J. Pharmacol.</i> , 457:137-146, 2002.
	C70	Dirnagl <i>et al.</i> , "Pathobiology of ischaemic stroke: an integrated view," <i>Trends Neurosci.</i> , 22:391-397, 1999.
	C71	Dixon <i>et al.</i> , "AICA-riboside: safety, tolerance, and pharmacokinetics of a novel adenosine-regulating agent," <i>J. Clin. Pharmacol.</i> , 31:342-347, 1991.
	C72	Duval, "Effects of statins on ischemic stroke: neuroprotection and/or triggering of apoptotic damage?" <i>Letters to the Editor</i> , p. 989-990.
	C73	Eisieik and Leijersfam, "The Inducible Form of Nitric Oxide Synthase (iNOS) in Insulin-Producing Cells," <i>Diabetes and Metabolism</i> , 20:116-122, 1994.
	C74	Evans <i>et al.</i> , "Differential effects of monoclonal antibodies to tumor necrosis factor alpha and gamma interferon on induction of hepatic nitric oxide synthase in experimental gram-negative sepsis," <i>Infect. Immun.</i> , 60:4133-4139, 1992.

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	C75	Faggioto and Paoletti, "Statins and blockers of the renin-angiotensin system: vascular protection beyond their primary mode of action," <i>Hypertension</i> , 34:987-996, 1999.
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	C78	Fenyk-Melody <i>et al.</i> , "Experimental autoimmune encephalomyelitis is exacerbated in mice lacking the <i>NOS2</i> gene," <i>J. Immunol.</i> , 160:2940-2946, 1998.
	C79	Fern, "Ischemia: astrocytes show their sensitive side," <i>Prog. Brain Res.</i> , 132:405-411, 2001.
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	C81	Forster <i>et al.</i> , "Inducible nitric oxide synthase expression in human cerebral infarcts," <i>Acta Neuropathol.</i> , 97:215-220, 1999.
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	C134	Iadecola <i>et al.</i> , "Inducible nitric oxide synthase gene expression in brain following cerebral ischemia," <i>J. Cereb. Blood Flow Metab.</i> , 15:378-384, 1995.
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	C142	Janero, "Nitric oxide (NO)-related pharmaceuticals: contemporary approaches to therapeutic NO modulation," <i>Free Radic. Biol. Med.</i> , 28:1495-1506, 2000.
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	C144	Johnson, "Multi-Purpose Medicine," <a href="http://my.abcnews.go.com">http://my.abcnews.go.com</a> 2000
	C145	Jord'heui <i>et al.</i> , "Oxidation and nitrosation of thiols at low micromolar exposure to nitric oxide. Evidence for a free radical mechanism," <i>J. Biol. Chem.</i> , 278:15720-15726, 2003.
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